

SEQUENCE LISTING

<110> Piddington, Christopher S.
 Petrie, Charles
 Shoemaker, Kimberly E.
 Bishop, Paul D.

<120> ZACE2: A HUMAN METALLOENZYME

<130> 99-24C1

<150> 60/133,952

<151> 1999-05-13

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<151> 1999-08-27

<150> 09/563,516

<151> 2000-05-03

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<170> FastSEQ for Windows Version 3.0

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1 5

ctt ctc agc ctt gtt gct gta act gct gct cag tcc acc att gag gaa Leu Leu Ser Leu Val Ala Val Thr Ala Ala Gln Ser Thr Ile Glu Glu 10 15 20 103

| _ | gcc Ala 25 | _ | | | _ | - | - | | | | _ | | ~ | - | - | 151 |
|---|-------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----|
| | tat Tyr | | | | | | | | | | | | | | | 199 |
| _ | gag Glu | | - | | | _ | | | _ | | _ | | _ | | | 247 |
| | tta Leu | _ | _ | _ | | | | - | | _ | | | | | _ | 295 |
| | cag Gln | | | | - | _ | | _ | - | - | - | | | | | 343 |
| | tct Ser 105 | | _ | | | _ | - | _ | _ | | | _ | | | | 391 |
| | aat Asn | | | | | | | | | | | | | | | 439 |
| - | aat Asn | | | - | - | | | | - | | | _ | | - | | 487 |
| | gca Ala | | | | | | | | | | | | | | | 535 |
| | aga Arg | | | - | | _ | _ | _ | | | | | - | | | 583 |
| | gtc Val 185 | - | | | | | | - | _ | | | | | _ | | 631 |

| | _ | | | _ | | | | | _ | | | | | ggc Gly | | • | 679 |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-------------------|-----|----------|-----|
| | | | | | | | | _ | _ | | - | | | ttt Phe 230 | _ | | 727 |
| | | | | | | _ | | | | _ | | - | | gca Ala | _ | | 775 |
| _ | _ | | - | | | | | | - | | | | _ | ctc Leu | | 8 | 823 |
| | | | | | | | | | | | | | | ctg Leu | | | 871 |
| | _ | | - | | | | _ | | | | | - | - | act Thr | _ | <u>(</u> | 919 |
| | | | | | | | | | | | | | | gag Glu 310 | | (| 967 |
| | | | | | | | | | | | _ | | | gga Gly | | 10 | 015 |
| | | | | | | | - | | | | - | _ | | gca Ala | - | 10 | 063 |
| | | | | | | | | | | | | | | atc Ile | | 13 | 111 |
| atg | tgc | aca | aag | gtg | aca | atg | gac | gac | ttc | ctg | aca | gct | cat | cat | gag | 1. | 159 |

| Met 360 | Cys | Thr | Lys | Val | Thr 365 | Met | Asp | Asp | Phe | Leu 370 | Thr | Ala | His | His | Glu 375 | |
|------------|-----|-----|-----|-----|-------------------|-----|-----|-----|-----|------------|-----|-----|-----|-----|------------|------|
| _ | | | | - | tat Tyr | - | _ | _ | | - | _ | | | | _ | 1207 |
| | _ | | | • | aat Asn | _ | | | | • | • | - | | - | | 1255 |
| - | | | | - | gcc Ala | | | _ | | | | | | | | 1303 |
| - | | | - | | caa Gln | - | _ | | - | | _ | | | | | 1351 |
| | | | | | acg Thr 445 | | | | | | | | | | | 1399 |
| | | | | | tgg Trp | | | | | | | | | | | 1447 |
| _ | | _ | | _ | tgg Trp | | | _ | _ | _ | _ | | _ | | | 1495 |
| | - | | - | | cat His | | | | | | | | | | | 1543 |
| | | - | | | gat Asp | | | | | | | | | | | 1591 |
| | | | | - | ttt Phe 525 | | | | | | | | | | | 1639 |

| gaa Glu | ggc Gly | cct Pro | ctg Leu | cac His 540 | aaa Lys | tgt Cys | gac Asp | atc Ile | tca Ser 545 | aac Asn | tct Ser | aca Thr | gaa Glu | gct Ala 550 | gga Gly | 1687 |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-----------------------|-------------------|-------------------|-------------------|-------------------|---------------------|------|
| cag Gln | aaa Lys | ctg Leu | ttc Phe 555 | aat Asn | atg Met | ctg Leu | agg Arg | ctt Leu 560 | gga Gly | aaa Lys | tca Ser | gaa Glu | ccc Pro 565 | tgg Trp | acc Thr | 1735 |
| cta Leu | gca Ala | ttg Leu 570 | gaa Glu | aat Asn | gtt Val | gta Val | gga Gly 575 | gca Ala | aag Lys | aac Asn | atg Met | aat Asn 580 | gta Val | agg Arg | cca Pro | 1783 |
| ctg Leu | ctc Leu 585 | aac Asn | tac Tyr | ttt Phe | gag Glu | ccc Pro 590 | tta Leu | ttt Phe | acc Thr | tgg Trp | ctg Leu 595 | aaa Lys | gac Asp | cag Gln | aac Asn | 1831 |
| aag | aat | tct | ttt | gtg | gga | tgg | agt | acc | gac | tgg | agt | сса | tat | gca | gac | 1879 |
| Lys 600 | Asn | Ser | Phe | Val | Gly 605 | Trp | Ser | Thr | Asp | Trp 610 | Ser | Pro | Tyr | Ala | Asp 615 | |
| caa G1n | agc Ser | atc Ile | aaa Lys | gtg Val 620 | Arg | ata Ile | agc Ser | cta Leu | aaa Lys 625 | Ser | gct Ala | ctt Leu | gga Gly | gat Asp 630 | aaa Lys | 1927 |
| gca Ala | tat Tyr | gaa Glu | tgg Trp 635 | Asn | gac Asp | aat Asn | gaa Glu | atg Met 640 | Tyr | ctg Leu | ttc Phe | cga Arg | tca Ser 645 | · Ser | gtt Val | 1975 |
| gca Ala | tat Tyr | gct Ala | Met | agg Arg | cag Gln | tac Tyr | ttt Phe 655 | . Leu | aaa Lys | gta Val | aaa Lys | aat Asr 660 | ı Glr | atg Met | att : Ile | 2023 |
| ctt Leu | ttt Phe | e Gly | gag Glu | gag Glu | gat Asp | gtg Val | Arg | gtg JVal | gct Alá | : aat a Asr | ttg Leu 675 | Lys | a cca s Pro | a aga o Arg | a atc g Ile | 2071 |
| tco Ser 680 | Phe | aat Asr | t tto n Phe | ttt Phe | gto Val 685 | Thr | gca `Ala | a cct a Pro | aaa Lys | a aat s Asr 690 | n Va | g tof I Ser | t gat ^ Asp | t ato | att e Ile 695 | 2119 |
| cc1 | t aga | a ac | t gaa | a gti | t gaa | a aag | g gco | c ato | c agg | g ato | g tco | c cg | g age | c cg | t atc | 2167 |

| Asn Asp Ala Phe Arg Leu Asn Asp Asn Ser Leu Glu Phe Leu Gly Ile 715 720 725 cag cca aca ctt gga cct cct aac cag ccc cct gtt tcc ata tgg ctg Gln Pro Thr Leu Gly Pro Pro Asn Gln Pro Pro Val Ser Ile Trp Leu 730 att gtt ttt gga gtt gtg atg gga gtg ata gtg gtt ggc att gtc atc Ile Val Phe Gly Val Val Met Gly Val Ile Val Gly Ile Val Ile 745 750 755 ctg atc ttc act ggg atc aga gat cgg aag aag aaa aat aaa gca aga Leu Ile Phe Thr Gly Ile Arg Asp Arg Lys Lys Lys Asn Lys Ala Arg 760 765 770 775 agt gga gaa aat cct tat gcc tcc atc gat att agc aaa gga gaa aat Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn 780 aat cca gga ttc caa aac act gat gat gtt caa acc tcc ttt Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe 795 800 tagaaaaaatc tatgttttc ctcttgaggt gattttgtt tatgtaaatg ttaattcat ggtatagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttca ggaaaaaaat tgtccaaaga caacatggcc aaggagagag catctcatt gacattgct tcagtattta tttctgtctc tggatttgac ttctgttctg | Pro Arg Thr Glu Val Glu Lys Ala Ile Arg Met Ser Arg Ser Arg Ile 700 705 710 | |
|---|--|--|
| att gtt ttt gga gtt gtg atg gga gtg ata gtg gtt ggc att gtc atc Ile Val Phe Gly Val Val Met Gly Val Ile Val Ile Val Gly Ile Val Ile 745 ctg atc ttc act ggg atc aga gat cgg aag aag aaa aat aaa gca aga Leu Ile Phe Thr Gly Ile Arg Asp Arg Lys Lys Lys Asn Lys Ala Arg 760 765 agt gga gaa aat cct tat gcc tcc atc gat att agc aaa gga gaa aat Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn 780 780 aat cca gga ttc caa aac act gat gat gtt cag aat cca gga ttc caa aac act gat gat gtt cag aat cca gga ttc caa aac act gat gat gtt cag ggataagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttca ggataagaaa ttcttgtttc tctcttgaggt gattttgttg tcagtattta tttctgtctc tggatttgac tctgttctg ttcagtattta tttctgtctc tggatttgac ttctgttctg | Asn Asp Ala Phe Arg Leu Asn Asp Asn Ser Leu Glu Phe Leu Gly Ile | 2215 |
| ctg atc ttc act ggg atc aga gat cgg aag aag aaa aat aaa gca aga Leu Ile Phe Thr Gly Ile Arg Asp Arg Lys Lys Lys Asn Lys Ala Arg 760 765 770 775 agt gga gaa aat cct tat gcc tcc atc gat att agc aaa gga gga aat Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn 780 785 790 aat cca gga ttc caa aac act gat gat gtt cag acc tcc ttt Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe 795 800 805 tagaaaaatc tatgttttc ctcttgaggt gattttgttg tatgtaaatg ttaatttcat ggtatagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttca ggaaaaaaat tytccaaaga caacatggcc aaggagaga catcttcatt gacattgt ttcagtatta tttctgtctc tggattgac ttctgttctg | Gln Pro Thr Leu Gly Pro Pro Asn Gln Pro Pro Val Ser Ile Trp Leu | 2263 |
| Leu Ile Phe Thr Gly Ile Arg Asp Arg Lys Lys Lys Asn Lys Ala Arg 760 765 770 775 agt gga gaa aat cct tat gcc tcc atc gat att agc aaa gga gaa aat 2407 Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn 780 785 790 aat cca gga ttc caa aac act gat gat gtt cag acc tcc ttt 2449 Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe 795 800 805 tagaaaaatc tatgttttc ctcttgaggt gattttgttg tatgtaaatg ttaattcat 2509 ggataagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgtca 2629 gaaaaaaaat tgtccaaaga caacatggcc aaggagagag catcttcatt gacattgctt 2629 tcagtattta tttctgtct tggatttgac ttctgttctg | Ile Val Phe Gly Val Val Met Gly Val Ile Val Val Gly Ile Val Ile | 2311 |
| Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn 780 785 790 aat cca gga ttc caa aac act gat gat gtt cag acc tcc ttt 2449 Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe 795 800 805 tagaaaaatc tatgttttc ctcttgaggt gattttgttg tatgtaaatg ttaattcat 2509 ggtatagaaa atataagatg ataaagatat cattaaatgt caaaaactatg actctgttca 2569 gaaaaaaaat tgtccaaaga caacatggcc aaggagagag catcttcatt gacattgctt 2629 tcagtattta tttctgtctc tggatttgac ttctgttctg | Leu Ile Phe Thr Gly Ile Arg Asp Arg Lys Lys Asn Lys Ala Arg | 2359 |
| Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe 795 800 805 tagaaaaatc tatgttttc ctcttgaggt gattttgttg tatgtaaatg ttaattcat 2509 ggtatagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttca 2569 gaaaaaaaat tgtccaaaga caacatggcc aaggagagag catcttcatt gacattgctt 2629 tcagtattta tttctgtctc tggatttgac ttctgttctg | Ser Gly Glu Asn Pro Tyr Ala Ser Ile Asp Ile Ser Lys Gly Glu Asn | 2407 |
| ggtatagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttca gaaaaaaaat tgtccaaaga caacatggcc aaggagagag catcttcatt gacattgctt 2629 tcagtattta tttctgtctc tggatttgac ttctgttctg | Asn Pro Gly Phe Gln Asn Thr Asp Asp Val Gln Thr Ser Phe | 2449 |
| agcagacact caataaatgc tagatttaca cactccttgt gctta 3334 | ggtatagaaa atataagatg ataaagatat cattaaatgt caaaactatg actctgttcagaaaaaaaat tgtccaaaga caacatggcc aaggagagag catcttcatt gacattgctt tcagtattta tttctgtctc tggatttgac ttctgttctg | 2569 2629 2689 2749 2809 2869 2929 2989 3049 3169 3169 |

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| Pro | Asn 290 | Ile | Asp | Val | Thr | Asp 295 | Ala | Met | Val | Asp | G1n 300 | Ala | Trp | Asp | Ala |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|--------------------|------------|------------|------------|------------|------------|
| G1n 305 | Arg | Ile | Phe | Lys | Glu 310 | Ala | Glu | Lys | Phe | Phe 315 | Val | Ser | Val | Gly | Leu 320 |
| Pro | Asn | Met | Thr | G1n 325 | Gly | Phe | Trp | Glu | Asn 330 | Ser | Met | Leu | Thr | Asp 335 | Pro |
| Gly | Asn | Val | G1n 340 | Lys | Ala | Val | Cys | His 345 | Pro | Thr | Ala | Trp | Asp 350 | Leu | Gly |
| Lys | Gly | Asp 355 | Phe | Arg | Ile | Leu | Met 360 | Cys | Thr | Lys | Val | Thr 365 | Met | Asp | Asp |
| | 370 | | | | | 375 | | | | | 380 | Tyr | | | |
| Tyr 385 | Ala | Ala | Gln | Pro | Phe 390 | Leu | Leu | Arg | Asn | G1 <i>y</i> 395 | Ala | Asn | Glu | Gly | Phe 400 |
| His | Glu | Ala | Val | G1y 405 | G1u | Пе | Met | Ser | Leu 410 | Ser | Ala | Ala | Thr | Pro 415 | Lys |
| | | | 420 | | | | | 425 | | | | Gln | 430 | | |
| G1u | Thr | G1u 435 | Ile | Asn | Phe | Leu | Leu 440 | Lys | Gln | Ala | Leu | Thr 445 | Ile | Val | Gly |
| | 450 | | | | , | 455 | | | - | • | 460 | Trp | | | |
| 465 | | | | | 470 | | | | | 475 | | Trp | | | 480 |
| | | | | 485 | | | | | 490 | | | His | | 495 | |
| | | • | 500 | | | | | 505 | | | | Asp | 510 | | |
| | | 515 | | | | | 520 | | | | | Phe 525 | | | |
| | 530 | | | | | 535 | | | | | 540 | Lys | | · | |
| 545 | | | | | 550 | | | | | 555 | | Met | | | 560 |
| - | - | | | 565 | · | | | | 570 | | | Val | | 575 | |
| | | | 580 | | | | | 585 | | | | Glu | 590 | | |
| | · | 595 | | | | | 600 | | | | | Gly 605 | | | |
| Asp | Trp 610 | Ser | Pro | Tyr | Ala | Asp 615 | Gln | Ser | Пе | Lys | Val 620 | Arg | Ile | Ser | Leu |

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Lys Ser Ala Leu Gly Asp Lys Ala Tyr Glu Trp Asn Asp Asn Glu Met
                    630
                                        635
Tyr Leu Phe Arg Ser Ser Val Ala Tyr Ala Met Arg Gln Tyr Phe Leu
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Lys Val Lys Asn Gln Met Ile Leu Phe Gly Glu Glu Asp Val Arg Val
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                                665
Ala Asn Leu Lys Pro Arg Ile Ser Phe Asn Phe Phe Val Thr Ala Pro
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                                                 685
Lys Asn Val Ser Asp Ile Ile Pro Arg Thr Glu Val Glu Lys Ala Ile
    690
                        695
                                             700
Arg Met Ser Arg Ser Arg Ile Asn Asp Ala Phe Arg Leu Asn Asp Asn
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Ser Leu Glu Phe Leu Gly Ile Gln Pro Thr Leu Gly Pro Pro Asn Gln
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                725
Pro Pro Val Ser Ile Trp Leu Ile Val Phe Gly Val Val Met Gly Val
            740
                                745
                                                     750
Ile Val Val Gly Ile Val Ile Leu Ile Phe Thr Gly Ile Arg Asp Arg
                            760
        755
                                                 765
Lys Lys Lys Asn Lys Ala Arg Ser Gly Glu Asn Pro Tyr Ala Ser Ile
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Asp Ile Ser Lys Gly Glu Asn Asn Pro Gly Phe Gln Asn Thr Asp Asp
785
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                                         795
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Val Gln Thr Ser Phe
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                                                                       120
taycarwsnw snytngcnws ntggaaytay aayacnaaya thacngarga raaygtncar
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aayatgaaya aygenggnga yaartggwsn genttyytna arqarcarws nacnytngen
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360

420

480 540

600

660

720

780 840

900

960

2220

2280

2340 2400

2415

300 caratgtayc cnytncarga rathcaraay ytnacngtna arytncaryt ncargcnytn carcaraayg gnwsnwsngt nytnwsngar gayaarwsna armgnytnaa yacnathytn aayacnatgw snacnathta ywsnacnggn aargtntgya ayccngayaa yccncargar tgyytnytny tngarccngg nytnaaygar athatggcna aywsnytnga ytayaaygar mgnytntggg cntgggarws ntggmgnwsn gargtnggna arcarytnmg nccnytntay gargartayg tngtnytnaa raaygaratg gcnmgngcna aycaytayga rgaytayggn gaytaytggm gnggngayta ygargtnaay ggngtngayg gntaygayta ywsnmgnggn carytnathg argaygtnga reayaentty gargaratha arcenytnta ygareayytn caygentayg tnmgngenaa rytnatgaay gentayeenw sntayathws neenathggn tgyytnccng cncayytnyt nggngayatg tggggnmgnt tytggacnaa yytntaywsn ytnacngtnc cnttyggnca raarccnaay athgaygtna cngaygcnat ggtngaycar gcntgggayg cncarmgnat httyaargar gcngaraart tyttygtnws ngtnggnytn ccnaayatga cncarggntt ytgggaraay wsnatgytna cngayccngg naaygtncar 1020 aargcngtnt gycayccnac ngcntgggay ytnggnaarg gngayttymg nathytnatg 1080 tgyacnaarg tnacnatgga ygayttyytn acngcncayc aygaratggg ncayathcar 1140 taygayatgg cntaygcngc ncarccntty ytnytnmgna ayggngcnaa ygarggntty 1200 caygargeng tnggngarat hatgwsnytn wsngengena encenaarea yytnaarwsn 1260 1320 athggnytny tnwsnccnga yttycargar gayaaygara cngarathaa yttyytnytn aarcargcny tnacnathgt nggnacnytn conttyacnt ayatgytnga raartggmgn 1380 tggatggtnt tyaarggnga rathccnaar gaycartgga tgaaraartg gtgggaratg 1440 aarmgngara thgtnggngt ngtngarccn gtnccncayg aygaracnta ytgygayccn 1500 gcnwsnytnt tycaygtnws naaygaytay wsnttyathm gntaytayac nmgnacnytn 1560 1620 taycarttyc arttycarga rgcnytntgy cargongona arcaygargg nccnytncay aartgygaya thwsnaayws nacngargen ggnearaary tnttyaayat gytnmgnytn 1680 ggnaarwsng arccntggac nytngcnytn garaaygtng tnggngcnaa raayatgaay 1740 gtnmgnccny tnytnaayta yttygarccn ytnttyacnt ggytnaarga ycaraayaar 1800 aaywsnttyg tnggntggws nacngaytgg wsnccntayg cngaycarws nathaargtn 1860 mgnathwsny tnaarwsngc nytnggngay aargcntayg artggaayga yaaygaratg 1920 tayytnttym gnwsnwsngt ngcntaygcn atgmgncart ayttyytnaa rgtnaaraay 1980 caratgathy tnttyggnga rgargaygtn mgngtngcna ayytnaarcc nmgnathwsn 2040 2100 ttyaayttyt tygtnacngc nccnaaraay gtnwsngaya thathccnmg nacngargtn garaargcna thmgnatgws nmgnwsnmgn athaaygayg cnttymgnyt naaygayaay 2160 wsnytngart tyytnggnat hcarccnacn ytnggnccnc cnaaycarcc nccngtnwsn athtggytna thgtnttygg ngtngtnatg ggngtnathg tngtnggnat hgtnathytn athttyacng gnathmgnga ymgnaaraar aaraayaarg cnmgnwsngg ngaraayccn taygcnwsna thgayathws naarggngar aayaayccng gnttycaraa yacngaygay gtncaracnw sntty

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<211> 16

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| tcc tgg ctc ctt ctc agc ctt gtt gct gtt act act gct cag tcc ctc Ser Trp Leu Leu Ser Leu Val Ala Val Thr Thr Ala Gln Ser Leu 5 10 15 20 | 165 |
| acc gag gaa aat gcc aag aca ttt tta aac aac ttt aat cag gaa gct Thr Glu Glu Asn Ala Lys Thr Phe Leu Asn Asn Phe Asn Gln Glu Ala 25 30 35 | 213 |
| gaa gac ctg tct tat caa agt tca ctt gct tct tgg aat tat aat act Glu Asp Leu Ser Tyr Gln Ser Ser Leu Ala Ser Trp Asn Tyr Asn Thr 40 45 50 | 261 |
| aac att act gaa gaa aat gcc caa aag atg agt gag gct gca gcc aaa Asn Ile Thr Glu Glu Asn Ala Gln Lys Met Ser Glu Ala Ala Ala Lys 55 60 65 | 309 |
| tgg tct gcc ttt tat gaa gaa cag tct aag act gcc caa agt ttc tca Trp Ser Ala Phe Tyr Glu Glu Gln Ser Lys Thr Ala Gln Ser Phe Ser 70 75 80 | 357 |
| cta caa gaa atc cag act ccg atc atc aag cgt caa cta cag gcc ctt | 405 |

| Leu 85 | Gln | Glu | Ile | Gln | Thr 90 | Pro | Ile | Ile | Lys | Arg 95 | G1n | Leu | Gln | Ala | Leu 100 | | |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-----------------------|-------------------|------------------------|---------------------|---|-----|
| cag Gln | caa Gln | agt Ser | ggg Gly | tct Ser 105 | tca Ser | gca Ala | ctc Leu | tca Ser | gca Ala 110 | gac Asp | aag Lys | aac Asn | aaa Lys | cag Gln 115 | ttg Leu | 2 | 453 |
| aac Asn | aca Thr | att Ile | ctg Leu 120 | aac Asn | acc Thr | atg Met | agc Ser | acc Thr 125 | att Ile | tac Tyr | agt Ser | act Thr | gga Gly 130 | aaa Lys | gtt Val | į | 501 |
| tgc Cys | aac Asn | cca Pro 135 | aag Lys | aac Asn | cca Pro | caa Gln | gaa Glu 140 | tgc Cys | tta Leu | tta Leu | ctt Leu | gag Glu 145 | cca Pro | gga Gly | ttg Leu | | 549 |
| gat Asp | gaa Glu 150 | ata Ile | atg Met | gcg Ala | aca Thr | agc Ser 155 | aca Thr | gac Asp | tac Tyr | aac Asn | tct Ser 160 | agg Arg | ctc Leu | tgg Trp | gca Ala | | 597 |
| tgg Trp 165 | gag Glu | ggc Gly | tgg Trp | agg Arg | gct Ala 170 | gag Glu | gtt Val | ggc Gly | aag Lys | cag Gln 175 | Leu | agg Arg | ccg Pro | ttg Leu | tat Tyr 180 | | 645 |
| gaa Glu | gag Glu | tat Tyr | gtg Val | gtc Val 185 | ctg Leu | aaa Lys | aac Asn | gag Glu | atg Met 190 | Ala | aga Arg | gca Ala | aac Asn | aat Asn 195 | tat Tyr | | 693 |
| aac Asn | gac Asp | tat Tyr | ggg Gly 200 | Asp | tat Tyr | tgg Trp | aga Arg | ggg Gly 205 | Asp | tat Tyr | gaa Glu | ıgca ıAla | gag Glu 210 | Gly | gca Ala | | 741 |
| gat Asp | ggc Gly | tac Tyr 215 | Asr | tat Tyr | aac Asn | cgt Arg | aac Asn 220 | Gln | ttg Leu | att Ile | gaa e G1ı | a gat u Asp 225 |) Val | gaa Glu | a cgt u Arg | | 789 |
| acc Thr | tto Phe 230 | e Alá | a gag a Glu | ato Ile | aag Lys | cca Pro 235 | Leu | tat Tyr | gag Glu | g cat u His | cti Lei 240 | His ل | gcc S Ala | tai Tyr | gtg Val | | 837 |
| agg Arg 249 | g Arg | aag Lys | g ttg s Lei | g atg u Met | gat Asp 25(| Thr | tac Tyr | cct Pro | tco Ser | tad Tyi 25 | ^ I](| c ago e Ser | c ccc r Pro | ac ⁻ Thi | t gga Gly 260 | | 885 |

| tgc (Cys l | ctc _eu | cct Pro | gcc Ala | cat His 265 | ttg Leu | ctt Leu | ggt Gly | gat Asp | atg Met 270 | tgg Trp | ggt Gly | aga Arg | ttt Phe | tgg Trp 275 | aca Thr | 933 |
|-------------------|-------------------|-------------------|-------------------|-----------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|
| aat d Asn l | ctg Leu | tac Tyr | cct Pro 280 | ttg Leu | act Thr | gtt Val | ccc Pro | ttt Phe 285 | gca Ala | cag Gln | aaa Lys | cca Pro | aac Asn 290 | ata Ile | gat Asp | 981 |
| gtt a | act Thr | gat Asp 295 | gca Ala | atg Met | atg Met | aat Asn | cag G1n 300 | ggc Gly | tgg Trp | gat Asp | gca Ala | gaa Glu 305 | agg Arg | ata Ile | ttt Phe | 1029 |
| caa (Gln | gag Glu 310 | gca Ala | gag Glu | aaa Lys | ttc Phe | ttt Phe 315 | gtt Val | tct Ser | gtt Val | ggc Gly | ctt Leu 320 | cct Pro | cat His | atg Met | act Thr | 1077 |
| caa G1n 325 | gga Gly | ttc Phe | tgg Trp | gca Ala | aac Asn 330 | tct Ser | atg Met | ctg Leu | act Thr | gag Glu 335 | cca Pro | gca Ala | gat Asp | ggc Gly | cgg Arg 340 | 1125 |
| aaa Lys | gtt Val | gtc Val | tgc Cys | cac His 345 | ccc Pro | aca Thr | gct Ala | tgg Trp | gat Asp 350 | ctg Leu | gga Gly | cac His | gga Gly | gac Asp 355 | ttc Phe | 1173 |
| aga Arg | atc Ile | aag Lys | atg Met 360 | tgt Cys | aca Thr | aag Lys | gtc Val | aca Thr 365 | Met | gac Asp | aac Asn | ttc Phe | ttg Leu 370 | Thr | gcc Ala | 1221 |
| cat His | cac His | gag Glu 375 | Met | gga Gly | cac His | atc Ile | caa Gln 380 | Tyr | gac Asp | atg Met | gca Ala | tat Tyr 385 | Ala | agg Arg | caa G1n | 1269 |
| cct Pro | ttc Phe 390 | Leu | cta Leu | aga Arg | aac Asn | gga Gly 395 | Ala | : aat ı Asn | gaa Glu | 999 Gly | ttc Phe 400 | His | gaa Glu | gct Ala | gtt Val | 1317 |
| gga Gly 405 | gaa Glu | ato Ile | atg Met | tca Ser | ctt Leu 410 | Ser | gca Ala | gct Ala | acc Thr | ccc Pro 415 |) Lys | cat His | ctg Leu | aaa Lys | tcc Ser 420 | 1365 |
| att Ile | ggt Gly | ctt Lei | t ctg . Lei | j cca i Pro 425 | Ser | : gat ` Asp | ttt Phe | caa e Glr | gaa Glu 430 | ı Asp | ago Ser | gaa Glu | aca ı Thr | gag Glu 435 | ata Ile | 1413 |

| aac ttc cta ctg Asn Phe Leu Leu 440 | ı Lys Gln Ala | | Val Gly Thr | | 1461 |
|---|---------------|-------------|-------------|-------------|------|
| act tac atg tta Thr Tyr Met Leu 455 | | | | | 1509 |
| ccc aaa gag cag Pro Lys Glu Glr 470 | | | | | 1557 |
| gtt ggt gtg gtg Val Gly Val Val 485 | | _ | - | | 1605 |
| gca tct ctg ttc Ala Ser Leu Phe | | | Ser Phe Ile | = | 1653 |
| aca agg acc att Thr Arg Thr Ile 520 | :Tyr Gln Phe | | Glu Ala Leu | · · | 1701 |
| gct aag tat aat Ala Lys Tyr Asr 535 | | | | | 1749 |
| gaa gct ggg cag Glu Ala Gly Glr 550 | | | | | 1797 |
| ccc tgg acc aaa Pro Trp Thr Lys 565 | | | | | 1845 |
| gta aaa cca ctg Val Lys Pro Leu | | • | Leu Phe Asp | | 1893 |
| gag cag aac aga | aat tct ttt | gtg ggg tgg | aac act gaa | tgg agc cca | 1941 |

| Glu | Gln | Asn | Arg 600 | Asn | Ser | Phe | Val | Gly 605 | Trp | Asn | Thr | Glu | Trp 610 | Ser | Pro | |
|-----|-----|-----|------------|------------|-----|-----|-----|------------|------------|-----|-----|-----|------------|-------------------|-----|------|
| | | _ | | _ | | | | _ | | - | | | | gct Ala | | 1989 |
| | - | | _ | | _ | | | | | - | _ | | - | ttc Phe | _ | 2037 |
| | | - | - | | | | | | | | | | | aaa Lys | | 2085 |
| ~ | | _ | | | | | - | - | - | - | | - | _ | ttg Leu 675 | | 2133 |
| | • | • | | | | | | • | | | | | | gtg Val | | 2181 |
| _ | - | | | - | | | - | | | | | | | tct Ser | | 2229 |
| | _ | | | | _ | | | _ | | - | | | | gag Glu | | 2277 |
| | | | | | | | | | | | | | | gtc Val | | 2325 |
| ata | tgg | ctg | att | att | ttt | ggt | gtt | gtg | atg | gca | ctg | gta | gtg | gtt | ggc | 2373 |
| Пе | Trp | Leu | Ile | Ile 745 | Phe | Gly | Val | Val | Met 750 | Ala | Leu | Val | Val | Val 755 | Gly | |
| atc | atc | atc | ctg | att | gtc | act | ggg | atc | aaa | ggt | cga | aag | aag | aaa | aat | 2421 |

| Ile | Ile | Ile | Leu 760 | Ile | Val | Thr | Gly | Ile 765 | Lys | Gly | Arg | Lys | Lys 770 | Lys | Asn | |
|-------------------|-------------------|-------------------|--------------------|------------|------------|-------------------|-------------------|-------------------|-------------|-------------------|-------------------|-------------------|------------|------------|------------|--------------|
| gaa Glu | aca Thr | aaa Lys 775 | aga Arg | gaa Glu | gag Glu | aac Asn | cct Pro 780 | tat Tyr | gac Asp | tcg Ser | atg Met | gac Asp 785 | att Ile | gga Gly | aaa Lys | 2469 |
| gga Gly | gaa Glu 790 | agc Ser | aat Asn | gca Ala | gga Gly | ttc Phe 795 | caa Gln | aac Asn | agt Ser | gat Asp | gat Asp 800 | gct Ala | cag Gln | act Thr | tcc Ser | 2517 |
| ttt Phe 805 | tago | caaaç | gca d | ttgt | cato | it to | cctg | tatg [.] | t aa | atgc [.] | taac | ttca | atagt | ac | | 2570 |
| | aaata gtcca | | agagt | tatad | ca ca | atgto | catt | a gc | tatc | aaaa | cta [.] | tgato | ctg t | tcag | gtaaac | 2630 2638 |
| | <; <; | 212> 213> | 805 PRT Mous | se | | | | | | | | | | | | |
| Mo+ | | 400> Sar | | Ser | Trn | l eu | l eu | l eu | Ser | Leu | Val | Ala | Val | Thr | Thr | |
| 1 | | | | 5 | | | | | 10 | | | | | 15 | | |
| Ala | G1n | Ser | Leu 20 | Thr | Glu | Glu | Asn | Ala 25 | Lys | Thr | Phe | Leu | Asn 30 | Asn | Phe | |
| Asn | Gln | G1u 35 | | G1u | Asp | Leu | Ser 40 | | Gln | Ser | · Ser | Leu 45 | Ala | Ser | Trp | |
| Asn | | Asr | Thr | Asn | Ile | Thr 55 | Glu | Glu | ı Asr | ı Ala | Gln 60 | Lys | Met | Ser | Glu | |
| Ala | 50 Ala | ı Ala | Lys | Trp | Ser | | Phe | . Tyr | · Glu | ı Glu | | Ser | Lys | Thr | Ala | |
| 65 | | | | | 70 | | | | | 75 | | | | _ | 08 | |
| Gln | ı Ser | · Phe | e Ser | Leu 85 | Gln | Glu | ı Ile | e Glr | n Thr 90 | Pro |) Ile | e Ile | Lys | Arg 95 | Gin | |
| Leu | ı Glr | ı Ala | Leu 100 | Gln | Gln | Ser | · Gly | Ser 105 | Ser | ^ A1a | a Lei | l Ser | Ala 110 | Asp | Lys | |
| Asr | ı Lys | Glr 115 | n Leu | | Thr | · Ile | e Leu 120 | Asr | | · Met | t Ser | Thr 125 | · Ile | | Ser | |

| Thr | Gly 130 | Lys | Val | Cys | Asn | Pro 135 | Lys | Asn | Pro | Gln | Glu 140 | Cys | Leu | Leu | Leu |
|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|
| Glu 145 | Pro | Gly | Leu | Asp | Glu 150 | Ile | Met | Ala | Thr | Ser 155 | Thr | Asp | Tyr | Asn | Ser 160 |
| Arg | Leu | Trp | Ala | Trp 165 | Glu | Gly | Trp | Arg | Ala 170 | Glu | Val | Gly | Lys | Gln 175 | Leu |
| Arg | Pro | Leu | Tyr 180 | Glu | Glu | Tyr | Val | Val 185 | Leu | Lys | Asn | Glu | Met 190 | Ala | Arg |
| Ala | Asn | Asn 195 | Tyr | Asn | Asp | Tyr | Gly 200 | Asp | Tyr | Trp | Arg | Gly 205 | Asp | Tyr | Glu |
| Ala | Glu 210 | Gly | Ala | Asp | Gly | Tyr 215 | Asn | Tyr | Asn | Arg | Asn 220 | G1n | Leu | Ile | Glu |
| Asp 225 | Val | Glu | Arg | Thr | Phe 230 | Ala | Glu | Ile | Lys | Pro 235 | Leu | Tyr | Glu | His | Leu 240 |
| His | Ala | Tyr | Val | Arg 245 | Arg | Lys | Leu | Met | Asp 250 | Thr | Tyr | Pro | Ser | Tyr 255 | Пе |
| Ser | Pro | Thr | Gly 260 | Cys | Leu | Pro | Ala | His 265 | Leu | Leu | Gly | Asp | Met 270 | Trp | Gly |
| Arg | Phe | Trp 275 | Thr | Asn | Leu | Tyr | Pro 280 | Leu | Thr | Val | Pro | Phe 285 | Ala | Gln | Lys |
| Pro | Asn 290 | Ile | Asp | Val | Thr | Asp 295 | Ala | Met | Met | Asn | G1n 300 | Gly | Trp | Asp | Ala |
| G1u 305 | Arg | Ile | Phe | G1n | G1u 310 | Ala | Glu | Lys | Phe | Phe 315 | Val | Ser | Val | Gly | Leu 320 |
| Pro | His | Met | Thr | G1n 325 | Gly | Phe | Trp | Ala | Asn 330 | Ser | Met | Leu | Thr | G1u 335 | Pro |
| Ala | Asp | Gly | Arg 340 | Lys | Val | Val | Cys | His 345 | Pro | Thr | Ala | Trp | Asp 350 | Leu | Gly |
| His | Gly | Asp 355 | Phe | Arg | Ile | Lys | Met 360 | Cys | Thr | Lys | Val | Thr 365 | Met | Asp | Asn |
| Phe | Leu 370 | | Ala | | | | | - | | | G1n 380 | Tyr | Asp | Met | Ala |
| Tyr 385 | Ala | Arg | Gln | Pro | Phe 390 | Leu | Leu | Arg | Asn | G1y 395 | Ala | Asn | Glu | Gly | Phe 400 |
| His | Glu | Ala | Val | Gly 405 | Glu | Ile | Met | Ser | Leu 410 | Ser | Ala | Ala | Thr | Pro 415 | Lys |
| His | Leu | Lys | Ser 420 | Ile | Gly | Leu | Leu | Pro 425 | Ser | Asp | Phe | Gln | G1u 430 | Asp | Ser |
| Glu | Thr | G1u 435 | Ile | Asn | Phe | Leu | Leu 440 | Lys | Gln | Ala | Leu | Thr 445 | Ile | Val | Gly |
| Thr | Leu 450 | Pro | Phe | Thr | Tyr | Met 455 | Leu | Glu | Lys | Trp | Arg 460 | Trp | Met | Val | Phe |

| Arg 465 | G1y | Glu | Ile | | Lys 470 | Glu | Gln | Trp | | Lys 475 | Lys | Trp | Trp | Glu | Met 480 |
|------------|-----|------------|------------|-----|--------------|-----|------------|------------|------------|--------------|-----|------------|------------|------------|------------|
| Lys | Arg | Glu | Ile | | Gly | Val | Val | Glu | Pro 490 | Leu | Pro | His | Asp | G1u 495 | Thr |
| Tyr | Cys | Asp | Pro 500 | | Ser | Leu | | His 505 | Val | Ser | Asn | Asp | Tyr 510 | Ser | Phe |
| Пе | Arg | Tyr 515 | Tyr | Thr | Arg | Thr | Ile 520 | Tyr | | Phe | Gln | Phe 525 | Gln | Glu | Ala |
| | 530 | | | | Lys | 535 | | | | | 540 | | | | |
| 545 | | | | | Ala 550 | | | | | 555 | | | | | 560 |
| | | | | 565 | Trp | | | | 570 | | | | | 5/5 | |
| | | | 580 | | Lys | | | 585 | | | | | 590 | | |
| | | 595 | | | | | 600 | | | | | 605 | | | Thr |
| | 610 | | | | | 615 | | | | | 620 | | | | Leu |
| 625 | | | | | 630 | | | | | 635 | | | | | Met 640 |
| | | | | 645 | | | | | 650 | | | | | 655 | |
| | | | 660 | | | | | 665 | | | | | 670 | | Val |
| | | 675 | | | | | 680 | | | | | 685 | | | Pro |
| | 690 | | | | | 695 | | | | | 700 | | | | Ile |
| 705 | | | | | 710 | | | | | 715 | ı | | | | Asn 720 |
| | | | | 725 | • | | | | 730 |) | | | | 735 | |
| | | | 740 |) | | | | 745 |) | | | | 750 |) | Leu |
| | | 755 |) | | | | 760 |) | | | | 765 | 5 | | / Arg |
| | 770 |) | | | | 775 |) | | | | 780 |) | | | Met |
| Asp 785 | | e Gly | / Lys | Gly | / Glu 790 | | · Asr | n Ala | a GIS | / Phe 795 | | ı ASY | ı ser | · AS | Asp 800 |

Ala Gln Thr Ser Phe 805 <210> 7 <211> 2415 <212> DNA <213> Artificial Sequence <220> <223> This degenerate sequence encodes the amino acid sequence of SEQ ID NO:6. <221> misc feature <222> (1)...(2415) <223> n = A.T.C or G<400> 7 60 atgwsnwsnw snwsntggyt nytnytnwsn ytngtngcng tnacnacngc ncarwsnytn acngargara aygcnaarac nttyytnaay aayttyaayc argargcnga rgayytnwsn 120 taycarwsnw snytngcnws ntggaaytay aayacnaaya thacngarga raaygcncar 180 aaratgwsng argengenge naartggwsn genttytayg argarearws naaraengen 240 carwsnttyw snythcarga rathcarach conathatha armgnoaryt noargonyth 300 carcarwsng gnwsnwsngc nytnwsngcn gayaaraaya arcarytnaa yacnathytn 360 420 aayacnatgw snacnathta ywsnacnggn aargtntgya ayccnaaraa yccncargar tgyytnytny tngarcengg nytngaygar athatggena enwsnaenga ytayaaywsn 480 mgnytntggg cntgggargg ntggmgngcn gargtnggna arcarytnmg nccnytntay 540 gargartayg tngtnytnaa raaygaratg genmgngena ayaaytayaa ygaytayggn 600 gaytaytggm gnggngayta ygargcngar ggngcngayg gntayaayta yaaymgnaay 660 carytnathg argaygtnga rmgnacntty gengaratha arcenytnta ygarcayytn 720 caygentayg tnmgnmgnaa rytnatggay aentayeenw sntayathws neenaenggn 780 840 tgyytnccng cncayytnyt nggngayatg tggggnmgnt tytggacnaa yytntayccn ytnacngtnc cnttygcnca raarccnaay athgaygtna cngaygcnat gatgaaycar 900 ggntgggayg cngarmgnat httycargar gcngaraart tyttygtnws ngtnggnytn 960 cencayatga encarggntt ytgggenaay wsnatgytna engareenge ngayggnmgn 1020 aargtngtnt gycayccnac ngcntgggay ytnggncayg gnqayttymg nathaaratg 1080 tgyacnaarg tnacnatgga yaayttyytn acngcncayc aygaratggg ncayathcar 1140 taygayatgg cntaygcnmg ncarcentty ytnytnmgna ayggngcnaa ygarggntty 1200 caygargeng tnggngarat hatgwsnytn wsngengena encenaarea yytnaarwsn 1260 athggnytny tnccnwsnga yttycargar gaywsngara cngarathaa yttyytnytn 1320 aarcargeny tnacnathgt nggnaenyth centtyaent ayatqytnga raartggmgn 1380

tggatgqtnt tymgnggnga rathccnaar garcartgga tgaaraartg gtgggaratg

aarmgngara thgtnggngt ngtngarccn ytnccncayg aygaracnta ytqygayccn

1440

1500

| gcnwsnytnt tycaygtnws naaygaytay wsnttyathm gntaytayac nmgnacnath taycarttyc arttycarga rgcnytntgy cargcngcna artayaaygg nwsnytncay aartgygaya thwsnaayws nacngargcn ggncaraary tnytnaarat gytnwsnytn ggnaaywsng arccntggac naargcnytn garaaygtng tnggngcnmg naayatggay gtnaarccny tnytnaayta yttycarccn ytnttygayt ggytnaarga rcaraaymgn aaywsnttyg tnggntggaa yacngartgg wsnccntayg cngaycarws nathaargtn mgnathwsny tnaarwsngc nytnggngcn aaygcntayg artggacnaa yaaygaratg ttyytnttym gnwsnwsngt ngcntaygcn atgmgnaart ayttywsnat hathaaraay caracngtnc cnttyytnga rgargaygtn mgngtnwsng ayytnaarcc nmgngtnwsn tytayttyt tygtnacnws nccncaraay gtnwsngayg tnathccnmg nwsngargtn gargaygcna thmgnatgws nmgnggnmgn athaaygayg tnttyggnyt naaygayaay wsnytngart tyytnggnat hcayccnacn ytngarccnc cntaycarcc nccngtnacn athtggytna thathttygg ngtngtnatg gcnytngtng tngtnggnat hathathytn athgtnacng gnathaargg nmgnaaraar aaraaygara cnaarmgnga rgaraayccn taygaywsna tggayathgg naarggngar wsnaaygcng gnttycaraa ywsngaygay gcncaracnw sntty | 1560 1620 1680 1740 1800 1860 1920 1980 2040 2160 2220 2280 2340 2415 |
|---|--|
| <210> 8 <211> 2638 <212> DNA <213> Mouse <220> <221> CDS | |
| <pre><222> (106)(2520) <400> 8 agtgcccaac ccaagttcaa aggctgatga gagagaaaaa ctcatgaaga gattttactc tagggaaagt tgctcagtgg atgggatctt ggcgcacggg gaaag atg tcc agc tcc</pre> | 60 117 |
| tcc tgg ctc ctt ctc agc ctt gtt gct gtt act act gct cag tcc ctc Ser Trp Leu Leu Ser Leu Val Ala Val Thr Thr Ala Gln Ser Leu 5 10 15 20 | 165 |
| acc gag gaa aat gcc aag aca ttt tta aac aac ttt aat cag gag gct Thr Glu Glu Asn Ala Lys Thr Phe Leu Asn Asn Phe Asn Gln Glu Ala 25 30 35 | 213 |
| gaa gac ctg tct tat caa agt tca ctt gct tct tgg aat tat aat act Glu Asp Leu Ser Tyr Gln Ser Ser Leu Ala Ser Trp Asn Tyr Asn Thr 40 45 50 | 261 |

| | | act Thr 55 | _ | - | | _ | | _ | _ | _ | _ | - | _ | _ | | 309 |
|-----|-----|-------------------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| | | gcc Ala | | | - | _ | _ | | _ | | - | | | | | 357 |
| | | gaa Glu | | _ | | _ | | | _ | - | | | _ | _ | | 405 |
| _ | | agt Ser | | | | - | | | _ | - | _ | | | _ | - | 453 |
| | | att Ile | _ | | | _ | _ | | | | _ | | | | _ | 501 |
| - | | cca Pro 135 | | | | | - | _ | | | | | | | _ | 549 |
| - | - | ata Ile | _ | | | | | | | | | | | | | 597 |
| | | ggc Gly | | | - | | _ | | - | _ | _ | | - | _ | | 645 |
| | | tat Tyr | | | | | | | | | | | | | | 693 |
| | | tat Tyr | | | | | | | | | | | | | | 741 |
| gat | ggc | tac | aac | tat | aac | cgt | aac | cag | ttg | att | gaa | gat | gta | gaa | cgt | 789 |

| Asp | Gly | Tyr 215 | Asn | Tyr | Asn | Arg | Asn 220 | Gln | Leu | Ile | Glu | Asp 225 | Val | Glu | Arg | | |
|-------------------|-------------------|-------------------|-----------------------|------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|------|---|
| acc Thr | ttc Phe 230 | gca Ala | gag Glu | atc Ile | aag Lys | cca Pro 235 | ttg Leu | tat Tyr | gag Glu | cat His | ctt Leu 240 | cat His | gcc Ala | tat Tyr | gtg Val | 837 | |
| agg Arg 245 | agg Arg | aag Lys | ttg Leu | atg Met | gat Asp 250 | acc Thr | tac Tyr | cct Pro | tcc Ser | tac Tyr 255 | atc Ile | agc Ser | ccc Pro | act Thr | gga Gly 260 | 885 | |
| tgc | ctc | cct | gcc | cat | ttg | ctt | ggt | gat | atg | tgg | ggt | aga | ttt | tgg | aca | 933 | |
| Cys | Leu | Pro | Ala | His 265 | Leu | Leu | Gly | Asp | Met 270 | Trp | Gly | Arg | Phe | Trp 275 | Thr | | |
| aat Asn | ctg Leu | tac Tyr | cct Pro 280 | ttg Leu | act Thr | gtt Val | ccc Pro | ttt Phe 285 | gca Ala | cag Gln | aaa Lys | cca Pro | aac Asn 290 | ata Ile | gat Asp | 981 | |
| gtt Val | act Thr | gat Asp 295 | gca Ala | atg Met | atg Met | aat Asn | cag Gln 300 | Gly | tgg Trp | gat Asp | gca Ala | gaa Glu 305 | Arg | ata Ile | ttt Phe | 1029 | |
| caa Gln | gag Glu 310 | A1a | gag Glu | aaa Lys | ttc Phe | ttt Phe 315 | ۷a٦ | tct Ser | gtt Val | ggc Gly | ctt Leu 320 | Pro | cat His | atg Met | act Thr | 1077 | |
| caa G1n 325 | Gly | tto Phe | tgg Trp | gca Ala | aac Asn 330 | Ser | atg Met | ctg : Leu | act Thr | gag Glu 335 | ı Pro | gca Ala | gat Asp | ggc Gly | cgg Arg 340 | 1125 |) |
| aaa Lys | gtt Val | gto Val | tgc Cys | cac His | Pro | aca Thr | gct Ala | tgg Trp | gat Asp 350 | Leu | gga Gly | a cac His | gga Gly | gac Asp 355 | ttc Phe | 1173 | } |
| aga Arg | ato Ile | aag Lys | g atg s Met 360 | : Cys | aca Thr | aag Lys | g gto Val | aca Thr 365 | Met | gad Asp | aac Asr | tto Phe | tte Lei 370 | ı Thr | a gcc r'Ala | 1221 | - |
| cat | cac | gaç | g ato | g gga | a cac | ato | caa | a tat | gac | atç | g gca | a tai | t gco | c agg | g caa | 1269 | } |

| His | His | G1u 375 | Met | Gly | His | Ile | G1n 380 | Tyr | Asp | Met | Ala | Tyr 385 | Ala | Arg | Gln | |
|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-------------------|-----------------------|-------------------|-------------------|-------------------|-------------------|--------------------|-----------------------|-------------------|------|
| Pro | ttc Phe 390 | ctg Leu | cta Leu | aga Arg | aac Asn | gga Gly 395 | gcc Ala | aat Asn | gaa Glu | ggg Gly | ttc Phe 400 | cat His | gaa Glu | gct Ala | gtt Val | 1317 |
| gga Gly 405 | gaa Glu | atc Ile | atg Met | tca Ser | ctt Leu 410 | tct Ser | gca Ala | gct Ala | acc Thr | ccc Pro 415 | aag Lys | cat His | ctg Leu | aaa Lys | tcc Ser 420 | 1365 |
| att Ile | ggt Gly | ctt Leu | ctg Leu | cca Pro 425 | tcc Ser | gat Asp | ttt Phe | caa Gln | gaa Glu 430 | gat Asp | agc Ser | gaa Glu | aca Thr | gag Glu 435 | ata Ile | 1413 |
| aac | ttc | cta | ctg | aaa | cag | gca | ttg | aca | att | gtt | gga | aca | cta | ccg | ttt | 1461 |
| Asn | Phe | Leu | Leu 440 | Lys | Gln | Ala | Leu | Thr 445 | Ile | Val | Gly | Thr | Leu 450 | Pro | Phe | |
| act Thr | tac Tyr | atg Met 455 | Leu | gag Glu | aag Lys | tgg Trp | agg Arg 460 | Trp | atg Met | gtc Val | ttt Phe | cgg Arg 465 | Gly | gaa Glu | att Ile | 1509 |
| ccc Pro | aaa Lys 470 | Glu | cag Gln | tgg Trp | atg Met | aaa Lys 475 | Lys | tgg Trp | tgg Trp | gag Glu | atg Met 480 | . Lys | cgg Arg | gag Glu | atc Ile | 1557 |
| gtt Val 485 | Gly | gtg Val | gtg Val | gag Glu | cct Pro 490 | Leu | cct Pro | cgt Arg | gat Asp | gaa Glu 495 | ı Thr | tac Tyr | tgt Cys | gac Asp | cct Pro 500 | 1605 |
| gca Ala | ı tct ı Ser | t ctg Lei | g tto u Phe | cat His | : Val | tct Ser | aat Asr | gat n Asp | tac Tyr 510 | Ser | tto Phe | att lle | cga Arg | a tai J Tyr 519 | tac Tyr | 1653 |
| aca Thr | a agg r Arg | g aco g Thr | att 11e 520 | e Tyr | caa Glr | a tto n Phe | c cag e Glr | g ttt n Phe 529 | e Glr | a gaa n Glu | a gct u Ala | ctt a Lei | tgt LCys 530 | s Glr | a gca n Ala | 1701 |
| gct | aag | g ta [.] | t aaf | t ggt | t tct | c ct | g cad | c aaa | a tgi | t gad | c ato | c tca | a aaf | t tc | c act | 1749 |

| Ala | Lys | Tyr 535 | Asn | Gly | Ser | Leu | His 540 | Lys | Cys | Asp | Ile | Ser 545 | Asn | Ser | Thr | |
|-----|-------------------|------------|-----|-----|-----|-----|------------|-----|-----|-----|-----|------------|-----|-----|-----|------|
| - | gct Ala 550 | | _ | - | _ | | _ | - | - | _ | | | | | | 1797 |
| | tgg Trp | | - | - | | - | | | | - | | | | | | 1845 |
| - | aaa Lys | | _ | | | | | | - | | | | | | | 1893 |
| | cag Gln | | | | | | | | | | | - | | | | 1941 |
| tat | gcc | gac | caa | agc | att | aaa | gtg | agg | ata | agc | cta | aaa | tca | gct | ctt | 1989 |
| Tyr | Ala | Asp 615 | Gln | Ser | Ile | Lys | Val 620 | Arg | Ile | Ser | Leu | Lys 625 | Ser | Ala | Leu | |
| | gct Ala 630 | | _ | | - | | | | | _ | _ | | _ | | - | 2037 |
| | tct Ser | - | - | | - | _ | _ | _ | | | | | | | | 2085 |
| | aca Thr | - | | | | | - | - | _ | - | | - | - | - | | 2133 |
| | aga Arg | _ | | | | | | _ | | | | | | | | 2181 |
| gat | gtc | att | cct | aga | agt | gaa | gtt | gaa | gat | gcc | atc | agg | atg | tct | cgg | 2229 |

| Asp Va | l Ile 695 | | Arg | Ser | Glu | Val 700 | Glu | Asp | Ala | Ile | Arg 705 | Met | Ser | Arg | |
|---------------------------|----------------------------------|-------------------|-------------------|-------|------------|------------|-------|-------|-------|------------|------------|-------------------|------|--------|--------------|
| ggc cgc Gly Arg 710 | ı Ile | | | - | | | - | | _ | | _ | | | | 2277 |
| ctg ggg Leu Gly 725 | • | | | | | | | | | _ | | | - | | 2325 |
| ata tgq Ile Trp | | | | | | _ | | | - | - | | | | | 2373 |
| atc atc | | _ | | - | | | | | | | | | | | 2421 |
| gaa aca Glu Thi | | Arg | _ | | | | | - | • | _ | _ | | | | 2469 |
| gga ga | a agc | aat | gca | gga | ttc | caa | aac | agt | gat | gat | gct | cag | act | tcc | 2517 |
| Gly Glu 79 | | Asn | Ala | Gly | Phe 795 | Gln | Asn | Ser | Asp | Asp 800 | Ala | Gln | Thr | Ser | |
| ttt tag Phe 805 | gcaaa | gca (| cttg [.] | tcato | et to | cctg1 | tatgi | t aaa | atgci | taac | ttc | atag [.] | tac | | 2570 |
| acaaaa gttgtc | | agag [.] | tatad | ca ca | atgto | catta | a gct | tatca | aaaa | ctat | tgato | ctg ⁻ | ttca | gtaaac | 2630 2638 |
| • | <210> <211> <212> <213> | 805 PRT | se | | | | | | | | | | | | |

<400> 9 Met Ser Ser Ser Trp Leu Leu Leu Ser Leu Val Ala Val Thr Thr Ala Gln Ser Leu Thr Glu Glu Asn Ala Lys Thr Phe Leu Asn Asn Phe Asn Gln Glu Ala Glu Asp Leu Ser Tyr Gln Ser Ser Leu Ala Ser Trp Asn Tyr Asn Thr Asn Ile Thr Glu Glu Asn Ala Gln Lys Met Ser Glu Ala Ala Ala Lys Trp Ser Ala Phe Tyr Glu Glu Gln Ser Lys Thr Ala Gln Ser Phe Ser Leu Gln Glu Ile Gln Thr Pro Ile Ile Lys Arg Gln Leu Gln Ala Leu Gln Gln Ser Gly Ser Ser Ala Leu Ser Ala Asp Lys Asn Lys Gln Leu Asn Thr Ile Leu Asn Thr Met Ser Thr Ile Tyr Ser Thr Gly Lys Val Cys Asn Pro Arg Asn Pro Gln Glu Cys Leu Leu Leu Glu Pro Gly Leu Asp Glu Ile Met Ala Thr Ser Thr Asp Tyr Asn Ser Arg Leu Trp Ala Trp Glu Gly Trp Arg Ala Glu Val Gly Lys Gln Leu Arg Pro Leu Tyr Glu Glu Tyr Val Val Leu Lys Asn Glu Met Ala Arg Ala Asn Asn Tyr Asn Asp Tyr Gly Asp Tyr Trp Arg Gly Asp Tyr Glu Ala Glu Gly Ala Asp Gly Tyr Asn Tyr Asn Arg Asn Gln Leu Ile Glu Asp Val Glu Arg Thr Phe Ala Glu Ile Lys Pro Leu Tyr Glu His Leu His Ala Tyr Val Arg Arg Lys Leu Met Asp Thr Tyr Pro Ser Tyr Ile Ser Pro Thr Gly Cys Leu Pro Ala His Leu Leu Gly Asp Met Trp Gly Arg Phe Trp Thr Asn Leu Tyr Pro Leu Thr Val Pro Phe Ala Gln Lys Pro Asn Ile Asp Val Thr Asp Ala Met Met Asn Gln Gly Trp Asp Ala Glu Arg Ile Phe Gln Glu Ala Glu Lys Phe Phe Val Ser Val Gly Leu

Pro His Met Thr Gln Gly Phe Trp Ala Asn Ser Met Leu Thr Glu Pro Ala Asp Gly Arg Lys Val Val Cys His Pro Thr Ala Trp Asp Leu Gly His Gly Asp Phe Arg Ile Lys Met Cys Thr Lys Val Thr Met Asp Asn Phe Leu Thr Ala His His Glu Met Gly His Ile Gln Tyr Asp Met Ala Tyr Ala Arg Gln Pro Phe Leu Leu Arg Asn Gly Ala Asn Glu Gly Phe His Glu Ala Val Gly Glu Ile Met Ser Leu Ser Ala Ala Thr Pro Lys His Leu Lys Ser Ile Gly Leu Leu Pro Ser Asp Phe Gln Glu Asp Ser Glu Thr Glu Ile Asn Phe Leu Leu Lys Gln Ala Leu Thr Ile Val Gly Thr Leu Pro Phe Thr Tyr Met Leu Glu Lys Trp Arg Trp Met Val Phe Arg Gly Glu Ile Pro Lys Glu Gln Trp Met Lys Lys Trp Trp Glu Met Lys Arg Glu Ile Val Gly Val Val Glu Pro Leu Pro Arg Asp Glu Thr Tyr Cys Asp Pro Ala Ser Leu Phe His Val Ser Asn Asp Tyr Ser Phe Ile Arg Tyr Tyr Thr Arg Thr Ile Tyr Gln Phe Gln Phe Gln Glu Ala Leu Cys Gln Ala Ala Lys Tyr Asn Gly Ser Leu His Lys Cys Asp Ile Ser Asn Ser Thr Glu Ala Gly Gln Lys Leu Leu Lys Met Leu Ser Leu Gly Asn Ser Glu Pro Trp Thr Glu Ala Leu Glu Asn Val Val Gly Ala Arg Asn Met Asp Val Lys Pro Leu Leu Asn Tyr Phe Gln Pro Leu Phe Asp Trp Leu Lys Glu Gln Asn Arg Asn Ser Phe Val Gly Trp Asn Thr Glu Trp Ser Pro Tyr Ala Asp Gln Ser Ile Lys Val Arg Ile Ser Leu Lys Ser Ala Leu Gly Ala Asn Ala Tyr Glu Trp Thr Asn Asn Glu Met Phe Leu Phe Arg Ser Ser Val Ala Tyr Ala Met Arg Lys Tyr Ser Ser

| I | 1e | Пe | Lys | Asn 660 | G1n | Thr | Val | Pro | Phe 665 | Leu | G1u | Glu | Asp | Va1 670 | Arg | Val | |
|---|-----------|------------|----------------|------------------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|------------|----|
| S | er | Asp | Leu 675 | | Pro | Arg | Val | Ser 680 | | Tyr | Phe | Phe | Val 685 | Thr | Ser | Pro | |
| G | 1n | Asn 690 | | Ser | Asp | Val | Ile 695 | | Arg | Ser | Glu | Val 700 | Glu | Asp | Ala | Ile | |
| | rg 05 | | Ser | Arg | Gly | Arg 710 | Ile | Asn | Asp | Val | Phe 715 | Gly | Leu | Asn | Asp | Asn 720 | |
| | | Leu | Glu | Phe | Leu 725 | Gly | Ile | His | Pro | Thr 730 | Leu | Glu | Pro | Pro | Tyr 735 | Gln | |
| Ρ | ro | Pro | Val | Thr 740 | Ile | Trp | Leu | Ile | I1e 745 | Phe | Gly | Val | Val | Met 750 | Ala | Leu | |
| ٧ | al | Val | Val 755 | | Пе | Ile | He | Leu 760 | Ile | Val | Thr | Gly | I1e 765 | | Gly | Arg | |
| L | .ys | Lys 770 | | Asn | Glu | Thr | Lys 775 | Arg | Glu | Glu | Asn | Pro 780 | Tyr | Asp | Ser | Met | |
| | sp '85 | Ile | Gly | Lys | Gly | G1u 790 | Ser | Asn | Ala | Gly | Phe 795 | | Asn | Ser | Asp | Asp 800 | |
| Д | \la | Gln | Thr | Ser | Phe 805 | | | | | | | | | | | | |
| | | < | :213> :220> | · 25 · DNA · Art | ific | | | ence | | | | | | | | | |
| | | < | <223> | · PCR | ≀ pri | mer. | | | | | | | | | | | |
| Ć | gac | | <400> jatc | | agcg | jtc a | ıacta | l | | | | | | | | | 25 |
| | | < | | > 20 > DNA | \ tific | cial | Sequ | ience | Ž | | | | | | | | |
| | | | <220> <223> | | R pri | imer. | | | | | | | | | | | |
| ı | aac | | <400> | | ncaat | .aa | | | | | | | | | | | 20 |